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☐ Expression of human brain carboxypeptidase B, a possible cleaving enzyme for beta-amyloid precursor protein, in peripheral fluids. *Neurosci Res.* 2001 Mar;39(3):313-7. PMID: 11248371 [PubMed - indexed for MEDLINE]

☐ 2: Matsumoto A, Itoh K, Matsumoto R. Related Articles, Links

☐ A novel carboxypeptidase B that processes native beta-amyloid precursor protein is present in human hippocampus. *Eur J Neurosci.* 2000 Jan;12(1):227-38. PMID: 10651877 [PubMed - indexed for MEDLINE]

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L2	1	brain near4(carboxypeptidase adj B)	USPAT	OR	OFF	2005/05/17 20:24
L3	51	human near4(carboxypeptidase adj B)	USPAT	OR	OFF	2005/05/17 20:24
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NEWS	17	APR 25	Patent searching, including current-awareness
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=> s l1 and APP
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DUPLICATE 1

AN 2002405327 EMBASE

TI **Brain-specific carboxypeptidase B**: Selective
down-regulation in ependymal cell by irradiation and altered
 β -amyloid processing.

AU Kawabe T.; Sasaki R.; Nishimura H.; Soejima T.; Matsuyama S.;
Sugimura K.;

Matsumoto A.

CS T. Kawabe, Division of Radiology, Kobe Univ. Grad. School of
Medicine,

Kusunoki-cho, Kobe 650-0017, Japan

SO Neuroscience Research Communications, (2002) Vol. 31, No. 2, pp.
75-84.

Refs: 23

ISSN: 0893-6609 CODEN: NRCOEE

CY United States

DT Journal; Article

FS 005 General Pathology and Pathological Anatomy

008 Neurology and Neurosurgery

014 Radiology

029 Clinical Biochemistry

LA English

SL English

ED Entered STN: 20021121

Last Updated on STN: 20021121

AB Human **brain carboxypeptidase B** (HBCPB) is a
brain protease expressed in a group of neuronal perikarya and
ependymal cells, and cleaves β -amyloid precursor protein (**APP**
) in vitro, generating β -amyloid ($A\beta$)-bearing C-terminal
fragment of **APP**. To explore its physiological function,
response to external stress, such as ionizing radiation, and
expression in

mouse brain were studied. Mouse head was irradiated at 10 Gy,
then brains

were excised before or after 1h until 9 d of irradiation. They
were

analyzed using TUNEL assay and anti C14-module antibody which
detects

epitopes unique to HBCPB, and western analysis. The mouse
HBCPB-homologue

cleaved mouse brain **APP** in vitro. Morphological study indicated
its identical distribution to HBCPB. During 2 h after
irradiation, the

HBCPB-homologue expression in edendymal cells, but not in neurons,
 decreases transiently, and recovers within 4 h. Concomitantly, processing of **APP** is altered, increasing generation of the C-terminal 12 kDa fragment. Since **APP** participates in cellular response to tolerance in CNS, this finding suggests that radiation-induced **APP** processing and differential response in its expression between ependymal cells and neurons are associated with important physiological function in CNS.

L3 ANSWER 2 OF 4 MEDLINE on STN DUPLICATE 2
 AN 2001339416 MEDLINE
 DN PubMed ID: 11248371
 TI Expression of human **brain carboxypeptidase B**
 , a possible cleaving enzyme for beta-amyloid precursor protein,
 in peripheral fluids.
 AU Matsumoto A; Motozaki K; Seki T; Sasaki R; Kawabe T
 CS Department of Radiation Biophysics and Genetics, Kobe University
 School of Medicine, Kusunoki-cho chuo-ki 7-5-1, Kobe650-0017, Japan...
 amat@med.kobe-u.ac.jp
 SO Neuroscience research, (2001 Mar) 39 (3) 313-7.
 Journal code: 8500749. ISSN: 0168-0102.
 CY Ireland
 DT Journal; Article; (JOURNAL ARTICLE)
 LA English
 FS Priority Journals
 EM 200106
 ED Entered STN: 20010618
 Last Updated on STN: 20010618
 Entered Medline: 20010614
 AB Human **brain carboxypeptidase B** (HBCPB) is a novel **brain** protease that processes native brain beta-amyloid precursor protein (**APP**) in vitro. Immunoblot analysis of human serum and cerebrospinal fluid (CSF) using anti C14-module antibody, which recognizes the C-terminal peptide unique to HBCPB, detected the 30 and 40 kDa immunoreactive bands. Analysis of HBCPB prepared from both serum and CSF demonstrated proteolytic activities for brain **APP**. Protease inhibitor spectrum analysis also supports that these bands correspond to the mature form and and prepro form of HBCPB, respectively. As is the case in brain parenchyma, the prepro-form is dominant in CSF. In serum,

however, the majority of HBCPB exists in the mature form, possibly due to an abundant trypsin-like proteolytic activity in serum. HBCPB expressed in serum and CSF, therefore, may have a significance as a peripheral marker of the brain protease, which participates in APP processing in human brain.

L3 ANSWER 3 OF 4 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation on STN

DUPLICATE 3

AN 2001:533671 BIOSIS

DN PREV200100533671

TI Distribution of a human **brain carboxypeptidase B** capable of cleaving beta-amyloid precursor protein (APP) in normal and Alzheimer's diseased brain.

AU Itoh, Kyoko [Reprint author]; Matsumoto, Akira

CS Department of Neurosciences, School of Medicine, Case Western Reserve

University, 2109 Adelbert Rd, Rm 658, Cleveland, OH, 44106-4975, USA

SO Acta Histochemica et Cytochemica, (2001) Vol. 34, No. 4, pp. 275-283.

print.

CODEN: ACHCBO. ISSN: 0044-5991.

DT Article

LA English

ED Entered STN: 14 Nov 2001

Last Updated on STN: 23 Feb 2002

AB The processing of beta-amyloid precursor protein (APP) is considered critical for understanding the pathogenesis of Alzheimer's disease (AD). To elucidate the significance of APP processing enzyme, we studied immunohistochemically the distribution of APP-processing protease (human carboxypeptidase B: HBCPB) in the normal control brains and AD brains, using anti-C14 antibody which recognizes C-terminal 14 amino acids of HBCPB. In the control brains, intense and diffuse C14-immunoreactivity was observed in the cytoplasm of pyramidal neurons of the hippocampus. Moderate immunoreactivity was found in the cortical and subcortical neurons. In AD brains, C14-immunoreactivity was markedly decreased in the brain regions examined except for the brain stem and cerebellum. However, HBCPB was shown to be colocalized with beta-amyloid protein (Abeta) in neuritic plaques. In addition, neuritic

plaques included C14-immunoreactive microglia/macrophages. Our present

studies indicate that the expression of a novel **APP**-processing protease is impaired in AD brains and may suggest the possible role of

HBCPB in the pathogenesis of AD.

L3 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2005 ACS on STN

AN 2000:790617 CAPLUS

DN 133:346507

TI Human **brain carboxypeptidase B**, cDNA, and use in drug screening and disease diagnosis

IN Matsumoto, Akira

PA Japan

SO PCT Int. Appl., 84 pp.

CODEN: PIXXD2

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.
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PI	WO 2000066717	A1	20001109	WO 2000-JP2878
20000501				

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CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
EP 1179588 A1 20020213 EP 2000-922936
20000501

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO

PRAI JP 1999-125169 A 19990430
WO 2000-JP2878 W 20000501

AB A novel human **brain carboxypeptidase B**

(CPB), its cDNA, recombinant expression, antibody, and use in screening of

ligands, activators, and inhibitors, are disclosed. Use as beta amyloid

(A β) production regulator, and therapeutic agent for diseases related to

A β accumulation in brain such as senile dementia, Alzheimer's disease, Down's syndrome, hereditary cerebral hemorrhage, head contusion,

is claimed. Diagnosis of and diagnostic reagent kits for those diseases

are also claimed. A novel carboxypeptidase B with amyloid precursor

protein (APP) cleaving activity was identified from human hippocampus and its cDNA was cloned by screening human hippocampus cDNA

library. Northern blot anal. for tissue distribution, Western blot anal.

with polyclonal antibodies, immunohistochem. studies, and homol. searches,

were carried out. Enzyme activity studies showed that A β 1-40 and

A β 1-42 were substrates.

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